Virginia Department of Health Brucellosis: Overview for Healthcare Providers

Organism	• Brucella spp. that infect humans: B. abortus, B. canis, B. ceti, B. inopinata, B. neotomae, B. melitensis,
	B. pinnepedalis, and B. suis
December 1 - D. Lillian	Brucella spp. are nonspore-forming, slow-growing, tiny, gram-negative coccobacilli.
Reporting to Public	• Suspected or confirmed cases require <u>immediate</u> notification to the local health department (LHD).
Health	See http://www.vdh.virginia.gov/local-health-districts/
Infectious Dose	• 10 to 100 organisms
Occurrence	 Worldwide, especially in Mediterranean Basin (Portugal, Spain, Southern France, Italy, Greece, Turkey, North Africa), Mexico, South and Central America, Eastern Europe, Asia, Africa, Caribbean and Middle East. ~115 cases in the United States and ~0-3 cases in Virginia are reported annually.
Natural Reservoir	 B. abortus (cattle), B. canis (dogs); B. ceti (dolphins, porpoises, whales); B. neotomae (wild rodents), B. melitensis (sheep, goats, camels), B. pinnipedialis (seals, sea lions, walruses), and B. suis (pigs). Infection might occur in other animals (e.g., bison, elk, coyotes, and some deer).
Route of Infection	 Multiple routes of infection, including ingestion of unpasteurized dairy products or undercooked meat from infected animals; inhalation of aerosols; contact through broken skin or mucous membranes with tissues, blood, urine, vaginal discharges, aborted fetuses and especially placentas of infected animals; inoculation with animal vaccine (injection or spraying into wounds or eyes); person-to-person transmission is rare, but can occur (see Communicability)
Communicability	Person-to-person transmission is rare, but has been reported with perinatal transmission, breastfeeding, blood transfusion or tissue transplantation, and sexual contact.
Risk factors	 Consuming unpasteurized dairy products, especially those from endemic areas; handling infected animals, tissues, or specimens during work (slaughterhouse workers, meat-packing workers, veterinarians, laboratory workers) or recreation (hunters of feral swine or other game animals)
Case-fatality Rate	• Low (< 1%), usually from endocarditis
Incubation Period	Highly variable, ranging from 5 days-6 months (average is 2-4 weeks)
Clinical Description	 Fever (constant or intermittent), chills, sweats, malaise, anorexia, headache, arthralgia, myalgia, back pain, fatigue, weight loss, depression, and pregnancy complications (spontaneous abortion, preterm delivery, miscarriage or intrauterine infection with fetal death). Musculoskeletal and genitourinary systems are commonly affected. Illness might last a few weeks to several months or longer.
Differential	Numerous because of nonspecific clinical presentation and varied complications
Diagnosis	
Radiography	Chest x-ray is often normal, but might show lung abscesses, single or miliary nodules, bronchopneumonia, enlarged hilar lymph nodes, or pleural effusions
Specimen	Alert lab if brucellosis is suspected so that appropriate precautions are taken during testing.
Collection and	Available tests are culture of blood, bone marrow or tissue; serology (EIA and agglutination tests); and
Laboratory Testing	PCR.
	 Brucella spp. are slow-growing bacteria. Culture might require extended incubation times (e.g., up to 21 days for non-automated broth blood culture and up to 10 days for automated systems). False-positive results with IgM EIA test can occur; further testing by agglutination test or testing paired sera to detect 4-fold change in titer is recommended. If brucellosis is suspected, notify LHD immediately to discuss the case. If VDH approves public health testing based on clinical presentation and exposure history, specimens may be sent to the Division of Consolidated Laboratory Services (DCLS). For questions about specimen collection, contact the DCLS Emergency Officer available 24/7 at 804-335-4617.
Treatment: Adults,	Combination therapy to decrease the incidence of relapse:
children ≥ 8 years*†	• Oral doxycycline (2–4 mg/kg per day, maximum 200 mg/day, in 2 divided doses) or oral tetracycline (30–40 mg/kg per day, maximum 2 g/day, in 4 divided doses) - and -

Infection Control	Use standard precautions for all patients.
	recommended. Serological monitoring is not available for <i>B. abortus</i> RB51 vaccine, but a baseline serum sample can be collected to rule out infection with other <i>Brucella</i> spp. if needed.
	Regular symptom watch (e.g., weekly) and daily self-fever checks through 24 weeks post-exposure are
	• For PEP regimens, see above and note that rifampin should not be used for RB51 vaccine exposures.
	exposure.
	• Self-inoculation with vaccine has occurred in veterinarians. Vaccine exposures typically occur through direct contact and individuals exposed to RB51 vaccine should be considered as having a high-risk
Vaccine*	• In the United States, a modified live vaccine (<i>B. abortus</i> RB51 vaccine) is licensed only for animals.
	(baseline), 6, 12, 18 and 24 weeks post-exposure is recommended. Serologic monitoring is not currently available for <i>B. abortus</i> RB51 vaccine or <i>B. canis</i> exposures.
	• For those with low- or high-risk exposures, regular symptom watch (e.g., weekly) and daily self-fever checks through 24 weeks post-exposure is recommended; sequential serological monitoring at 0
	appropriate antimicrobial, should be considered. Pregnant women should consult their obstetrician.
	• For patients with contraindications to doxycycline or rifampin, TMP-SMZ, in addition to another
. ,	 Doxycycline (100 mg twice daily for 3 weeks) plus rifampin (600 mg once daily for 3 weeks).
Prophylaxis*	exposures, but can be considered on a case-by-case basis.
Postexposure	 sulfamethoxazole, if doxycycline is not used) and rifampin for a minimum of 6 weeks. PEP is recommended for high-risk exposures; PEP is generally not recommended for low-risk
	included for the first 7 to 14 days of therapy, in addition to doxycycline (or trimethoprim-
	complications, such as deep tissue abscesses. Red Book® recommends a 3-drug regimen: gentamicin
	• Notes: Case-fatality rate is < 1%. Surgical intervention should be considered in patients with
etc.)*	extended for 4 to 6 months.
osteomyelitis,	• For life-threatening complications, such as meningitis or endocarditis, duration of therapy often is
meningitis,	 Rifampin can be used in combination with this regimen to decrease the rate of relapse.
Complicated Cases (endocarditis,	(or TMP-SMZ if tetracyclines are contraindicated). Streptomycin might not be readily available in the United States.
Treatment:	• Streptomycin or gentamicin for the first 14 days of therapy in addition to a tetracycline for 6 weeks
Pregnancy*	antimicrobial therapy instructions.
Treatment:	• Tetracyclines are contraindicated for pregnant patients. Consult obstetrician regarding specific
	in 1 or 2 divided doses.
	of age. Red Book® (2018) lists dose for rifampin as 15–20 mg/kg per day, maximum 600–900 mg/day,
v o years	is contraindicated. Tetracyclines (such as doxycycline) should be avoided in children less than 8 years
Children < 8 years*	mg/kg per day, maximum 2.4 g/day) divided in 2 doses for 4 to 6 weeks. • Notes: For combination therapy, consider adding rifampin. Consult physician for dosing or if rifampin
Treatment:	• Oral TMP-SMZ (trimethoprim, 10 mg/kg per day, maximum 480 mg/day; and sulfamethoxazole, 50
	day, maximum 200 mg/day, in 2 divided doses.
	tetracyclines are contraindicated. Red Book® (2018) lists dose for oral doxycycline as 4.4 mg/kg per
	• Notes: Combination therapy with trimethoprim-sulfamethoxazole (TMP-SMZ) can be used if
	• Recommended for a minimum of 6 weeks.
	• Rifampin (15–20 mg/kg per day, maximum 600–900 mg/day, in 1 or 2 divided doses).

*Adapted from CDC Brucellosis Reference Guide: Exposures, Testing, and Prevention (February 2017) and Red Book®: 2018-2021 Report of the Committee on Infectious Diseases, 31st ed. Elk Grove Village, IL: American Academy of Pediatrics;2018: 257. Publications cited by CDC for treatment recommendations: 1) Ariza J et al. 2007. Perspectives for the Treatment of Brucellosis in the 21st Century: The Ioannina Recommendations. PLoS Med. 4(12): e317.; 2) Al-Tawfiq JA. 2008. Therapeutic options for human brucellosis. Expert Rev Anti Infect Ther. 6(1): 109-120.; 3) Solera J. 2010. Update on brucellosis: therapeutic challenges. Intl J Antimicrob Agent. 36S, S18–S20. Note that the B. abortus strain used in the RB51 vaccine was derived by selection in rifampin-enriched media and is resistant to rifampin in vitro. This strain is also resistant to penicillin. If the infection is due to this vaccine strain, treatment should be determined accordingly (for example, doxycycline and TMP-SMZ in place of rifampin). Specifics on the treatment or PEP regimen and dose should be established in consultation with the person's healthcare provider in case of contraindications to the aforementioned. For additional information on dosing, please consult with the package inserts.

[†] VDH modified this category to include children aged 8 years or older based on communication with CDC.